



IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

APPLICANT:

Wieder

SERIAL NO.:

10/521,841

EXAMINER: Meera Natarajan, PhD

601-1-134PCTUS

FILED:

July 27, 2005

ART UNIT:

1643

FOR

ALPHA 5 BETA 1 AND ITS ABILITY TO REGULATE THE CELL

SURVIVAL PATHWAY

DECLARATION OF ROBERT WIEDER

- I, Robert Wieder, hereby declare that:
- 1. I am a citizen of the United States and reside at 1380 Outlook Drive West, Mountainside, NJ 07092.
- 2. I received a BA degree in Biochemistry and Biology from the University of Pennsylvania in 1976, a Master of Philosophy (M. Phil.) and a Ph.D. in Biomedical Sciences from the Mount Sinai School of Medicine of the City University of New York in 1982 and an M.D. degree from the Mount Sinai School of Medicine in 1983. I trained in Internal Medicine at the Montefiore Medical Center of the Albert Einstein School of Medicine from 1983 to 1985, as a Medical Staff Fellow at the National Heart, Lung and Blood Institute of the National Institutes of Health from 1985 to 1988, and as a fellow in Medical Oncology/Hematology at Memorial Sloan Kettering Cancer Center from 1988 to 1992. I was employed as an attending physician on the Autologous Bone Marrow Transplant Service at Memorial Sloan Kettering Cancer Center from 1992 to 1993. I have been employed at the University of Medicine and Dentistry of New Jersey-New Jersey Medical School as a faculty member from 1993 to the present time. The details of my education and professional history are set forth in my curriculum vitae, attached hereto as Exhibit A.
 - 3. I have over 33 years experience in the fields of biomedical sciences and medicine.

4. I am the author or co-author of more than 36 scientific articles on the subjects of biology, biophysics and oncology. A list of these articles is set forth in my curriculum vitae, attached hereto. My current area of research involves dormancy and survival signaling in breast cancer micrometastases in the bone marrow and interaction of micrometastatic breast cancer cells with the bone marrow microenvironment.

5. I am an inventor of the subject matter disclosed and claimed in United States Patent Application Number 10/521,841 (hereinafter the '841 application).

Statements Regarding the Defects of Varner and Li et al.

- 6. I have read and discussed with counsel the Official Action dated April 24, 2009, received in connection with the '841 application. I understand the nature of the rejections made by the Examiner concerning Varner [United States Patent Number (USPN) 7,311,911] in view of Li et al. (United States Publication Serial Number 2004/0048312).
- 7. The Office Action acknowledges that Varner does not teach or suggest adjuvant therapy. As an expert in the field, I can confirm that there is no teaching in Varner that relates to an adjuvant setting. Indeed, as taught throughout Varner, the intended therapeutic purpose with regard to patients with cancer is to reduce or inhibit tumor angiogenesis. It is well understood in the field of clinical oncology that a therapeutic purpose directed to reducing or inhibiting tumor angiogenesis only pertains to a patient with detectable disease (i.e., a detectable tumor or tumors). In contrast, my invention relates to a method for treating a patient in an adjuvant setting, wherein the patient fails to exhibit signs of detectable disease. It is, therefore, apparent that the method of the present invention and that of Varner are directed to distinct patient populations.
 - 8. In light of the above, the statements in the Office Action pertaining to Varner

allegedly teaching that administration to a subject can either be over a relatively short period of time or can be over a more prolonged period of time and that different therapeutic protocols can be used to achieve the most effective regimen are misapplied. In keeping with the teachings of Varner, such guidance only relates to a method for reducing or inhibiting angiogenesis for therapeutic purposes. Wherein the therapeutic purpose relates to the treatment of cancer in a patient, the intended purpose is to reduce or inhibit tumor angiogenesis. Thus, the teaching of Varner pertaining to treatment duration and therapeutic protocol would be understood by an ordinarily skilled practitioner to apply to treatment of a patient with detectable disease.

9. The Office Action looks to Li et al. for purportedly teaching adjuvant therapy. The Office Action concludes that the combined teachings of Varner and Li et al. would allegedly lead an ordinarily skilled practitioner to realize my invention. I disagree for a variety of reasons. To begin, the Li et al. patent application is directed in its entirety to integrin av \(\beta \) and its role in human cancers and a monoclonal antibody (mBLA3) specific for integrin avβ6. As shown in Figure 1 of Exhibit B (attached), I assessed and did not observe an increase in the expression of either αv or β6 in the dormant (growth arrested) breast cancer cells analyzed as taught in the present specification. For the record, Figure 1 of Exhibit B is identical to that of Figure 5A or 5B of the instant specification (United States Publication Number 2006/0035825), except that the arrows have been altered to be directed to the positions corresponding to αv and β6 mRNA on the microarray. Furthermore, Figure 13A of the instant specification demonstrates clearly and unambiguously that adhesion of dormant growth arrested breast cancer cells to fibronectin, a ligand for both integrins $\alpha 5\beta 1$ and $\alpha \nu \beta 6$, depends on integrin $\alpha 5$ but is unaffected by blocking antibody to integrin av. This complements the gene array expression data demonstrating a lack of expression of integrin ανβ6 in dormant breast cancer cells with a functional assay demonstrating a dependence of survival on integrin a5, which is contrasted with a complete lack of dependence on integrin av. These results demonstrate unequivocally that av and \(\text{\theta} \) do not play a significant role in the survival of dormant (growth arrested) breast cancer cells in the bone marrow microenvironment.

10. Moreover, in view of the teaching in the instant specification regarding the identity of the microarray used for the analysis presented in Figures 5A and 5B (see, for example, paragraph [0031]), an ordinarily skilled practitioner would have been able to identify which spots on the grid correspond to αv and $\beta 6$ mRNA on the microarray should such a practitioner have been motivated to investigate the expression of these mRNAs in the context of dormant breast cancer cell micrometastases. Indeed, I perform similar evaluations on microarray analyses published in the literature as a matter of routine practice. In short, all that an ordinarily skilled practitioner requires in this regard is the supplier and product name/catalog number of the microarray, because the identity of the particular positions/spots on the array is publicly available. Thus, in the unlikely event that an ordinarily skilled practitioner might have thought to combine the teachings of Varner and Li et al., the microarray data (Figures 5A and 5B) and the functional data (Figure 13A) presented in the instant specification would have discouraged such a practitioner from pursuing the combination. My results suggest that such a combination of teachings would not have had a reasonable expectation for success because $\alpha v \beta 6$ integrin is not expressed in dormant breast cancer cell micrometastases

11. The results presented in Figure 1 and general knowledge in the field also underscore the fact that integrins represent a very diverse family of proteins. Integrins exhibit dissimilar functions and such functions are, in turn, modified differentially in the context of different cellular environments. Thus, an ordinarily skilled practitioner would appreciate that what is determined to be true for one integrin cannot accurately be extrapolated to apply to any other integrin. That being said, there is no scientific basis for imagining that teaching the use of antibodies to a particular integrin in an adjuvant setting extends to the use of antibodies to any other integrin in any adjuvant setting. An ordinarily skilled practitioner would appreciate that in the absence of evidence indicating which integrin or integrins are relevant in a particular clinical setting with respect to expression and activity, there is no way to predict whether antibodies to a particular integrin will have any clinical utility. Furthermore, my data demonstrate that antibodies to $\alpha v\beta 6$ integrin are unlikely to have any clinical utility in the context of breast cancer

cell micrometastases in the bone marrow because $\alpha \nu \beta 6$ integrin is not expressed on these cells. See Exhibit B.

- 12. Moreover, the literature appears to show a consensus that $\alpha\nu\beta6$ increases with carcinogenesis in most primary tumors and some metastases, but primarily in squamous cell carcinomas, as well as in colon and ovarian epithelial cells. Expression appears to increase with cell crowding. There are no data on expression in breast cancer metastases. In the paper by Van Aarsen et al. (submitted previously), only a minority of primary breast carcinomas express this fibronectin binding integrin. Accordingly, since there are no data on metastasis, no significant role attributed to primary breast tumors, no evidence of expression in metastatic breast cancer cells and no evidence of an increase in dormant cells, there appears to be no prior data supporting a role for integrin $\alpha\nu\beta6$ in survival of dormant micrometastatic breast cancer cells in the bone marrow. The literature, therefore, fails to provide support for a role for integrin $\alpha\nu\beta6$ as a target for adjuvant therapy.
- 13. In summary, Varner teaches therapeutic regimens that inhibit angiogenesis and call for inhibition of $\alpha5\beta1$ integrin signaling, but Varner fails to teach adjuvant therapy. The Li et al. application mentions adjuvant therapy in passing, but is focused in its entirety on $\alpha\nu\beta6$ integrin and its role in human cancers and a monoclonal antibody (mBLA3) specific for $\alpha\nu\beta6$ integrin. An ordinarily skilled practitioner would appreciate that $\alpha5\beta1$ integrin and $\alpha\nu\beta6$ integrin are functionally disparate integrins and would not, therefore, consider it obvious to apply teachings pertaining to one integrin to that of the other. The literature, moreover, fails to teach or suggest any role for $\alpha\nu\beta6$ integrin in growth arrested cancer cells in general or growth arrested breast cancer cells in particular, nor any evident role in metastatic or micrometastatic breast cancer. In contrast, the literature affirms a role for $\alpha\nu\beta6$ integrin in rapidly proliferating cancer cells (i.e., non-dormant cancer cells). My data demonstrate that $\alpha\nu\beta6$ integrin is not expressed in dormant (growth arrested) breast cancer cells in the bone marrow microenvironment. See Figures 5A, 5B, and 13A of the application as filed and Figure 1 of Exhibit B. Taken together, these facts

would dissuade an ordinarily skilled artisan from combining the teachings of Varner and Li et al. Indeed, the facts in combination teach away from the presently claimed invention.

14. I further declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code, and that such willful statements may jeopardize the validity of the above-referenced application or any patent issued thereon.

RUBERT WIEDER, MO,

NAME

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EXHIBIT A

CURRICULUM VITAE

NAME:

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Robert Wieder

HOME ADDRESS:

1380 Outlook Drive West

Telephone: 908-377-1465

Telephone: (973) 972-4871

E-mail: wiederro@umdnj.edu

Mountainside, New Jersey 07092

OFFICE ADDRESS:

UMDNJ-New Jersey Medical School

Cancer Center H1216

185 South Orange Avenue

Newark, NJ 07103

1. EDUCATION:

<u>a.</u> <u>Undergraduate</u>:

University of Pennsylvania, Philadelphia, PA 1976 B.A. in Biochemistry and Biology

b. Graduate and Professional:

Mount Sinai School of Medicine of the City University of NY, New York, NY

1982 M.Phil. in Biomedical Sciences

1982 Ph.D. in Biomedical Sciences

1983 M.D.

2. POSTDOCTORAL TRAINING

<u>a.</u> <u>Internship and Residencies:</u>

1983-1985 Intern and Junior Resident, Internal Medicine, Montefiore Hospital Medical Center, Albert Einstein College of Medicine, Bronx, NY

1985-1986 Medical Staff Fellow, Clinical Associate, National Heart, Lung and Blood

Institute, NIH, Bethesda, MD

1988-1992 Fellow, Medical Oncology/Hematology, Memorial Sloan-Kettering

Cancer Center, New York, NY

b. Research Fellowships:

1986-1988 Medical Staff Fellow, Laboratory of Molecular Hematology,

National Heart, Lung and Blood Institute, NIH, Bethesda, MD

3. MILITARY

None

4. LICENSURE

<u>a.</u> N.J. Medical License: MA 60226, 1993

b. Other State Medical License: N.Y., 159293, 1984

5. CERTIFICATION

Board Certified:

Internal Medicine, #107202

Medical Oncology, #107202

6. NARCOTICS CERTIFICATION

U.S. BW 3903261 N.J. DO 63847

NJ Medical School

7. UNIVERSITY APPOINTMENTS

1992-1993	Instructor, Cornell University Medical College, New York, NY
1992-1993	Instructor, Department of Medicine, Memorial Sloan-Kettering
	Cancer Center, New York, NY
1993-2000	Assistant Professor, Department of Medicine,
	UMDNJ-New Jersey Medical School
1997-presen	t Assistant Professor, Department of Microbiology and Molecular
	Genetics, UMDNJ-New Jersey Medical School
1997-presen	t Assistant Professor, Department of Microbiology and Molecular
	Genetics, UMDNJ-Graduate School of Biomedical Sciences
1996-1999	Interim Director, Division of Oncology, Department of Medicine,
	UMDNJ-New Jersey Medical School
1999-2001	Interim Director, Division of Oncology/Hematology, Department of
	Medicine, UMDNJ-New Jersey Medical School
1998-2001	Associate Director, Clinical Research, UMDNJ-Cancer Center/ Newark
•	t Research Associate Member, Cancer Institute of New Jersey
•	t Associate Professor, Dept. of Medicine, UMDNJ-NJ Medical School
2005-presen	t Director, Clinical Research Office, UMDNJ-NJ Medical
	School/University Hospital Cancer Center

8. HOSPITAL APPOINTMENTS

1983-1985	Montenore Hospital, Medical Center, Bronx, NY, Housestaff
1983-1985	North Central Bronx Hospital, Bronx, NY, Housestaff
1985-1988	NIH Clinical Center, Medical Staff Fellow
1988-1992	Memorial Sloan-Kettering Cancer Center, New York, NY, Fellow,
	Medical Oncology/Hematology
1988-1992	Cornell Medical Center, New York, NY, Fellow, Medical
	Oncology/Hematology
1992-1993	Memorial Sloan-Kettering Cancer Center, Attending Physician Leukemia
	Service, Autologous Bone Marrow Transplant Unit
1994-present	UMDNJ-University Hospital, Medical Oncology/Hematology

2009-present Co-Medical Director, Center for Clinical and Translational Science, UMDNJ-

9. OTHER PROFESSIONAL POSITIONS AND MAJOR VISITING APPOINTMENTS

July-Sept., 1987 Special Fellow, Adult Bone Marrow Unit, Memorial Sloan-Kettering Cancer Center, New York, NY

10. AWARDS AND HONORS

1976	B.A. Cum Laude and Honors in Biochemistry
1980	NIH Medical Student Research Fellow
1981-1982	Mount Sinai Medical Scientist Fellow
1988-1989	American Cancer Society Clinical Oncology Fellow
1990-1991	Mortimer J. Lacher Fellow
1990-1992	Memorial Sloan-Kettering Cancer Center Clinical
	Scholars Biomedical Research Fellow
1990-1992	Charles A. Dana Fellow
1992	ASCO Travel Award
1994	Foundation of the University of Medicine and
	Dentistry of New Jersey Grant Award
1994	US Army Breast Cancer Research Prog. Career Development Award
1998	State of New Jersey Commission on Cancer Research Outstanding
	Breast Cancer Researcher Award
2001	State of New Jersey Commission on Cancer Research Service
	Recognition Award
2001	UMDNJ-New Jersey Medical School Faculty Association Clinical
	Sciences Faculty of the Year

11. BOARDS OF DIRECTORS/TRUSTEES none

12. MAJOR COMMITTEE POSITIONS AND MAJOR VISITING APPOINTMENTS

Extrainstitutional

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	1995-1996	Member, Special Awards Committee, American Soc. Clinical Oncology
	1995-pres.	NJ State Comm. on Cancer Research Breast Cancer Advisory Board, Chair 1998-2001, 2006-present
	1996-2005	Program Review Committee, Annual New Jersey Breast Cancer
		Research Symposium
	1997	Program Review Committee, Annual Scientific Retreat of the Cancer
		Institute of New Jersey and The New Jersey State Commission on
		Cancer Research
	1997-2004	Cancer Institute of New Jersey Protocol Advisory Committee
	1999	Research Planning Committee, Annual Scientific Retreat of the Cancer
		Institute of New Jersey and The New Jersey State Commission on
		Cancer Research
	1999-2000	Member, State of New Jersey Department of Health and Senior Services
		<u>Circle of Friends</u> Advisory Board
	2002-present	Extrainstitutional Member, Mount Sinai School of Medicine Institutional
		Biosafety Committee
	2003	Program Review Committee, Annual Scientific Retreat of the Cancer

Institute of New Jersey and The New Jersey State Commission on Cancer Research

2009-present Cancer Institute of New Jersey Clinical Trial Network Steering Committee

Institutional

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NIH	

1886-1988 NIH Bioethics Liaison Group

1987 Consultant, Working Group on General Information, Human Gene

Therapy Subcommittee, NIH Recombinant DNA Advisory Committee.

<u>UMDNJ</u>

1993-1997 Medical School Admissions Committee

1994-present M.D./Ph.D Committee, UMDNJ-NJMS/Graduate School Biomedical

Sciences, Member Program Committee and Advisory Council, Associate

Director (Clinical) - 1998-2003

1994-1995 Dean's Search Committee for Chair of Biochemistry and Molecular

Biology, New Jersey Medical School

1994-1995 Search Committee for Director of Oncology/Hematology, UMDNJ-

NJ Medical School

1993-1997 NJMS Research Office Summer Student Research Program Advisory

Comm.

1993-1996 NJMS Dept. of Medicine Committee on Medicare Patient Outcomes

Research

1995-present NJMS Faculty Investigator Group

1995 Search Committee for Director of Oncology, Beth Israel Medical Center,

Newark, NJ.

1995-1997 Dean's Biomedical Research Support Committee

1995-2001 Founding Member Internal Medicine Resident Research Committee

1995-1996 Dean's Self-Study Task Force, Med. School Clin. Science Depts.

Commitee

1995-2000 Screening Access of Value to Elderly Women Coalition (SAVE),

Women's Wellness Center, UMDNJ-New Jersey Medical School

1996-2003 UMDNJ/University Hospital Oncology Committee

1996 Dean's Committee on the Oncology Program

1996-2000 Dean's Research Advisory Group

1996-2000 IAMS Research Planning Committee

1996-2000 UMDNJ Committee on Bloodless Surgery and Medicine

1997-1998 Internal Medicine Residency Curriculum Committee

1997-1998 Dean's Committee on Research Space Allocation at NJ Medical School

1997-1998 Steering Committee, the Cancer Center at New Jersey Medical School

1998 Dean's Space Advisory Committee

1998-2000 NJMS Research Planning and Priorities Committee

1998-1999 Cancer Center Executive Committee

1999-2001 NJMS cancer education program committee

1999-2001 UMDNJ-UH Pain Management Committee

1999-2000 UMDNJ-UH Cancer Program Survey Team 2000

2000-2001 Cancer Center Steering Committee

2001	Ad hoc Searle, Pew and Sinsheimer Scholars review committees
2001	UMDNJ-NJMS Dept. Medicine Retreat Research Planning Committee
2001-2003	UMDNJ-NJMS Department of Medicine Research Committee
2001-2004	Dean's Search Committee for Chair of Pediatrics, NJ Medical School
2002-2006	UMDNJ-NJMS Cancer Center Animal Facility Advisory Committee
2003-2007	UMDNJ-NJMS Basic Science/Translational Task Force
2003-present	t UMDNJ-Newark Campus Institutional Review Board
2004-2005	UMDNJ Research Conflicts of Interest Committee
2004	NJMS/UH Cancer Center Director Search Committee
2004-presen	t NJMS Cancer Education Program Executive Committee
2004-presen	t NJMS Biomedical Research Support Committee
2004-presen	t NJMS-UH Cancer Center Committee for Basic and Translational
	Research
2004-2005	UMDNJ-NJMS Search Committee for Faculty of Biostatistics
2005	UMDNJ-NJMS/UH Cancer Center Search Committee for Chief
	Operating Officer
2005-presen	tNJMS-UH Cancer Center Faculty Search Committee
2007	Coordinator of NJMS-UH Cancer Center Science Grand Rounds
2007-2008	UMDNJ-NJMS Faculty Committee on Appointments and Promotions
2009-presen	tMember, Clinical Research Leadership Group, UMDNJ-NJMS Center
	for Clinical and Translational Science

13. <u>MEMBERSHIPS, OFFICES AND COMMITTEE ASSIGNMENTS IN PROFESSIONAL</u> SOCIETIES

- 1992 Member, The American Society of Hematology
- 1994 Member, American Society of Clinical Oncology; Special Awards Committee (95-96)
- 1994 Member, American Association for Cancer Research
- 1995 Member, American Society for Blood and Marrow Transplantation
- 1997 Member, The Harvey Society
- 2002 Member, European Society of Medical Oncology
- 2005 Member, Metastasis Research Society

14. MAJOR RESEARCH INTERESTS

- a. Dormancy and survival signaling in breast cancer
- b. Roles of retinoids and vitamin D analogues in breast cancer
- c. Gene therapy for treatment of cancer
- d. Clinical trials in Oncology in minority patients

15. GRANT HISTORY

Past support

a. Principal investigator

Title: Clinical Oncology Fellowship #88-144.

Agency: American Cancer Society

Direct Costs: \$10,000. Period: 7/1/88-6/30/89 Title: Training Grant #T32 CA-09512-07

Agency: NIH

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Direct Costs: \$30,000. Period: 7/1/89-6/30/90

Title: Biomedical Scholar Award

Agency: Memorial Sloan-Kettering Cancer Center

Direct Costs: \$60,000. Period: 7/1/90-6/30/92

Title: Leukemia Research, unrestricted

Agency: United Food and Commercial Workers Union Leukemia Fund.

Direct Costs: \$5,000.

Period: 1995

Title: Apoptosis in breast cancer, unrestricted

Agency: Friends of Charity Direct Costs: \$ 15,500.

Period: 1995

Title: Breast Cancer Research Symposium of New Jersey Agency; New Jersey Commission on Cancer Research

Direct Costs: \$10,000.

Period: 1996

Title: Research grant: The Role of basic fibroblast growth factor in human breast cancer.

Agency: Foundation of the University of Medicine and Dentistry of New Jersey.

Direct Costs: \$ 48,900.

Period: 7/1/94-6/30/95 (extended without additional funds to 6/30/97)

Title: Career Development Award AIBS #200, DAMD17-94-J-4463: The Role of basic fibroblast growth factor in human breast cancer. Agency: DOD, U.S. Army Breast Cancer Research Program.

Direct Costs: \$ 200,000. Period: 10/1/94-9/30/98

Title: Phase II study of the efficacy of Doxil (Doxorubicin HCl liposome injection, Sequus

Pharmaceuticals Inc., Menlo Park, CA 94025) in stage IV breast cancer.

Agency: Seguus Pharmaceuticals, Inc., Menlo Park, CA

Direct Costs: \$ 14,100. Period: 1997-1999

Title: Phase I/II trial of 13-cis-retinoic acid (Accutane), Paclitaxel (Taxol) and Carboplatin in

recurrent of metastatic squamous cell carcinoma of the head and neck.

Agency: Bristol-Myers, Inc. Direct Costs: \$ 46,059. Period: 1997-1999 Title: Differentiation of Breast Cancer by Retinoids and Vitamin D₃ Agency: State of New Jersey Commission on Cancer Research

Direct Costs: \$ 45,455. Period: 1998-2000

Title: A Phase I Study of Oral ILX 23-7553 Administered Daily × 5 Every 2 Weeks in Patients

With Solid Tumors Agency: Ilex Oncology Direct Costs: \$ 52,800. Period: 1999-2002

Title: Roles of all-trans retinoic acid and vitamin D₃ in potentiating cell death signaling by

Taxotere in breast and prostate cancer

Agency: Aventis, Inc. Direct Costs: \$ 133,500. Period: 2000-2002

Title: Potentiation of Taxotere-induced cytotoxicity by flavopiridol in breast cancer cells

Agency: Aventis, Inc. Direct Costs: \$ 98,464. Period: 2002-2003

Title: Research grant: DAMD17-01-C-0343:

The Roles of FGF-2, TGF Beta, and TGF Beta Receptor 2 in Breast Cancer Dormancy.

Agency: DOD, U.S. Army Breast Cancer Research Program.

Direct Costs: \$ 230,509. Period: 7/1/01-6/30/03

Title: The Role of Bone Marrow Stromal FGF-2 in Breast Cancer Dormancy.

Agency: State of New Jersey Commission on Cancer Research 02-1140-CCR-E0 Direct Costs: \$ 136,364. (Collaboration with Rider U.) Direct to our lab: \$68,182.

Period: 6/1/02-5/30/04

Title: Phase II Trial of 13-cis retinoic acid (Accutane), paclitaxel (Taxol) and paraplatin (Carboplatin) in invasive, recurrent or metastatic squamous cell carcinoma of the cervix

Agency: Bristol-Myers, Inc. Direct Costs: \$ 122,946. Period: 2001-2004

Title: Research grant: DAMD17-03-1-0524:

Overcoming Bone Marrow Stroma-Mediated Chemoresistance in Metastatic Breast Cancer

Cells.

Agency: DOD, U.S. Army Breast Cancer Research Program.

Direct Costs: \$ 312,903. Period: 7/1/03-6/30/06

Title: Phase II trial of Vesanoid (TRETINOIN, all-trans retinoic acid) and Taxol (Paclitaxel) in

patients with stage IV breast cancer

Agency: Bristol-Myers, Inc. Direct Costs: \$ 70,000. Period: 1999-2006

Title: Mechanisms involved in the treatment and prevention of breast cancer by 1,25-

dihydroxyvitaminD₃

Co-equal PI: with Sylvia Christakos, Department of Biochemistry and Molecular Biology,

UMDNJ-NJMS

Agency: NJMS-UH Cancer Center Research Grants Program

Direct Costs: \$80,000. Period: 2006-2007

Title: Signal pathway activation signature of cisplatin resistance in head and neck cancer

Role: Co-equal PI (with Erik Cohen) 20% effort

Agency: Foundation of UMDNJ

Direct Costs: \$ 70,000. Period: 2007-2008

Title: Effects of perscription coverage on control of cancer pain: 08-1096-CCR-EO

Role: PI 10% effort

Agency: New Jersey Commission on Cancer Research

Direct Costs: \$ 18,000. Period: 2007-2008

Title: Elimination of dormant breast cancer cells by targeting survival signaling

Role: Pl 10% effort

Agency: Ruth Estrin Goldberg Foundation

Direct Costs: \$ 25,000. Period: 2008-2009

b. Co-Investigator:

Title: Summer Fellowship (with Karen J. Finnigan): The roles of basic FGF in breast cancer

Agency: New Jersey Commission on Cancer Research

Direct Costs: \$2,800. Period: 7/1/94-8/31/94

Title: Outstanding Breast Cancer Research Fellowship (with Qin Wang): Modulation of

apoptosis by basic FGF in breast cancer

Agency: State of New Jersey Commission on Cancer Research

Direct Costs: \$50,000. Period: 9/5/97-9/4/99

Title: Post-Doctoral Supplement to NJCCR Breast Cancer Research Fellowship (Qin Wang)

Agency: Foundation of UMDNJ

Direct Costs: \$10,000. Period: 9/5/97-9/4/99

Title: Community Cancer Screening for Black/Hispanic Women

Agency: Centers for Disease Control and Prevention

Direct Costs: \$5,000. Period: 9/30/99-9/29/00

Title: Summer Fellowship (with Ilan Seth Weinberg): Eligibility screening for a Phase I colon

cancer vaccine protocol

Agency: New Jersey Commission on Cancer Research

Direct Costs: \$2,800. Period: June-August, 2000

Title: Summer Fellowship (with Jeaudine Evadne Bontemps): NSABP P-2 Trial: Study of

Tamoxifen and Raloxifene (STAR) for the prevention of breast cancer

Agency: New Jersey Commission on Cancer Research

Direct Costs: \$2,800. Period: June-August, 2000

Title: Summer Fellowship (with Michael Lindy): Overcoming Breast Cancer Dormancy in the

Bone Marrow Microenvironmnet

Agency: New Jersey Commission on Cancer Research

Direct Costs: \$2,800. Period: June-August, 2003

Title: Summer Fellowship: Mediation of Survival Signaling in Dormant Breast Cancer Cells

through PI3 kinase and Rho

PI: Bryan Benn

Agency: New Jersey Commission on Cancer Research

Direct Costs: \$2,800. Period: June-August, 2004

Title: Inhibition of Breast Cancer Growth by Vitamin D

PI: Sylvia Christakos, Department of Biochemistry and Molecular Biology, UMDNJ-NJMS

Agency: AHEPA Foundation

Dircet Costs: \$10,000. Period: 2004-2005

Title: Summer Fellowship (with Joanna Sesti): Role retinoic acid in disrupting survival

signaling in dormant breast cancer cells

Agency: New Jersey Commission on Cancer Research

Direct Costs: \$2,800. Period: June-August, 2006

Title: Potentiation of radiation-induced cytotoxicity in 1483 head and neck squamous cell

carcinoma by COX-2 inhibition

Co-PI: Erik G. Cohen, Department of Surgery, UMDNJ-NJMS

Agency: Foundation of UMDNJ

Direct Costs: \$50,000.

Period: 2004-2006

Title: Signal pathway activation signature of cisplatin resistance in head and neck cancer

Role: Joint Co-PI with Erik Cohen 20% effort

Agency: Foundation of UMDNJ

Direct Costs: \$ 70,000. Period: 2007-2008

Title: Role of Integrin Signaling in Resistance to Chemotherapy in Head and Neck

Squamous Cell Carcinoma

Role: Co- PI (with Erik Cohen) 10% effort Agency: Ruth Estrin Goldberg Foundation

Direct Costs: \$ 25,000. Period: 2007-2008

Present support

a. Principal investigator

Title: Use and evaluation of an ethnically-matched patient navigator to increase minority

patient recruitment to breast cancer clinical trials

Role: PI 10% effort

Agency: The Susan G. Komen Breast Cancer Foundation of Northern New Jersey

Direct Costs: \$195,000. Period: 2007-2010

Title: Cancer Center Patient Navigator Program

Role: PI 20% effort

Agency: C.R. Bard Foundation

Direct Costs: \$ 50,000. Period: 2007-2009

Title: Research grant: W81XWH-09-1-0119

Reactivation of Breast Cancer Micrometastases by Senescent Bone Marrow Stroma

(Human)

Agency: DOD, U.S. Army Breast Cancer Research Program.

Role: PI 15% effort Direct Costs: \$ 375,000. Period: 7/1/09-6/30/12

Title: Reactivation of breast cancer micrometastases by senescent bone marrow stroma

Role: PI 20%

Agency: NJMS Dean's Annual Bridge Grant Program

Direct Costs: \$25,000. Period: 2008-2010

Title: Research grant: 1 R21 CA142537-01A1

Reactivation of Breast Cancer Micrometastases by Senescent Bone Marrow Stroma

(Murine) Agency: NCI

Role: PI 11.6% effort Direct Costs: \$ 242,000. Period: 7/1/09-6/30/11

Title: Minority Breast Cancer Navigator Program

Role: PI 10%

Agency: Susan G. Komen for the Cure of Northern New Jersey

Direct Costs: 16,000. Period: 2009-2010

Title: UMDNJ-NJMS/UH Cancer Center Clinical Research Program

Role: PI 15%

Agency: New Jersey Commission on Cancer Research - New Jersey Cancer Research

Development Award Direct Costs: \$ 427,595. Period: 2009-2010

Title: Research grant: 1U10CA128506-01A1

Minority-Based CCOP at UMDNJ-NJ Medical School/University Hospital Cancer Center

Agency: NCI

Role: PI 30% effort Direct Costs: \$ 1,188,911. Period: 8/24/09-5/31/12

b. Co-investigator

Provisional Patent disclosures filed

2002 Alpha5 Beta1 and its Ability to Regulate the Cell Survival Pathway 60/396,482 Application filed 2003 - 601-1-134PCT

European Patent filing 7/16/03 Application/Patent no. 03799816.8-2403-US0321954

2003 Development of microfluidics technology to map cell surface protein signatures for use in laboratory investigations, diagnosis, prognosis and treatment of disease. – in collaboration with Sarnoff Corporation, Princeton, NJ

2004 Molecular and Cellular Retardation Methods and the Encounter, or Interaction, Complex. – in collaboration with Sarnoff Corporation, Princeton, NJ

16. MAJOR TEACHING EXPERIENCE

1975-76 Teaching Assistant, Organic Chemistry Laboratory, Department of Chemistry,

University of Pennsylvania, Philadelphia, PA.

1993-1998 Participated in Dept. Microbiology and Mol. Genetics Mol. Biology Journal Club

1993	Lectured in the New Jersey Medical School Immunology course
1994-1996	Supervised Clinical Oncology Fellows on the wards and in clinic
1994-1996	Conducted Clinical Oncology Journal Club
1994-preser	nt Lectured to New Jersey Medical School Internal Medicine Residents
1994-preser	nt Lectured annually in the New Jersey Medical School Introduction to Clinical Sciences course, Participated in physical diagnosis sessions
1994-preser	nt Lectured annually in Medical Student Summer Cancer Biology Research Program
1995, 1997	Lectured in the UMDNJ-Graduate School of Biomedical Sciences Biology of Human Tumors Course,
1995-2001	Lectured annually in the New Jersey Graduate School Microbial Genetics II Course
2001	UNDNJ-NJMS Mini Med School - How Cancer is Treated
2001, 2003,	2005 Lectured in the UMDNJ-Graduate School of Biomedical Sciences
	Molecular and Immunopathologic Mechanisms of Cancer
2003	Lectured in Oncology Nurses training course, UMDNJ-University Hospital
2003	Lectured in UMDNJ-NJMS Medical Residents Conference, UMDNJ-UH
2004	Lectured in UMDNJ-NJMS Physical Medicine and Rehabilitation Residents
	Conference, Kessler Institute for Rehabilitation
2006	Lectured in UMDNJ-NJMS Medical Residents Conference, UMDNJ-UH
2006	Medical Student Transition Curriculum presentation on CBCs
2008	Lectured in UMDNJ-NJMS Medical Residents Conference, UMDNJ-UH
2008, 2009	Course Director, Principles of Clinical and Translational Research in
	Oncology, UMDNJ-GSBS GSND 5235Q
2009	Lectured in UMDNJ-NJMS Medical Residents Conference, UMDNJ-UH

MEMBER GRADUATE THESIS OR EXAMINATION COMMITTEES

- Jennifer B. Jones (Department of Laboratory Medicine and Pathology, UMDNJ-Graduate School of Biomedical Sciences)
- Jackie Washington (Department of Microbiology and Molecular Genetics, UMDNJ-Graduate School of Biomedical Sciences)
- Daniel Aviv (Department of Microbiology and Molecular Genetics, UMDNJ-Graduate School of Biomedical Sciences)
- Xuening Wang (Department of Laboratory Medicine and Pathology, UMDNJ-Graduate School of Biomedical Sciences)
- Qing Mei Wang (Department of Laboratory Medicine and Pathology, UMDNJ-Graduate School of Biomedical Sciences)
- James Nugent (Department of Microbiology and Molecular Genetics, UMDNJ-Graduate School of Biomedical Sciences)
- Achal Trivedi (Department of Microbiology and Molecular Genetics, UMDNJ-Graduate School of Biomedical Sciences)
- Roman Wernyj (Department of Biochemistry and Molecular Biology, UMDNJ-Graduate School of Biomedical Sciences)
- Jennifer Czarneski (Department of Laboratory Medicine and Pathology, UMDNJ-Graduate School of Biomedical Sciences)

- Sahba Kianifard (Department of Microbiology and Molecular Genetics, UMDNJ-Graduate School of Biomedical Sciences)
- Melanie K. Lenahan (Department of Microbiology and Molecular Genetics, UMDNJ-Graduate School of Biomedical Sciences)
- Kathy Piparo (Department of Biochemistry and Molecular Biology, UMDNJ-Graduate School of Biomedical Sciences)
- Megan Fredericks (Department of Biochemistry and Molecular Biology, UMDNJ-Graduate School of Biomedical Sciences)
- Wei Bu (Department of Microbiology and Molecular Genetics, UMDNJ-Graduate School of Biomedical Sciences)
- Anoop Kavirayani (Department of Microbiology and Molecular Genetics, UMDNJ-Graduate School of Biomedical Sciences)
- Pedro L. Rodriguez (Biomedical Sciences Program, UMDNJ-Graduate School of Biomedical Sci.) Shan Jiang (Department of Biochemistry and Molecular Biology, UMDNJ-Graduate School of Biomedical Sciences)

Zhaoyu Sun (Biomedical Sciences Program, UMDNJ-Graduate School of Biomedical Sci.)
Gwen Mahon (Biomedical Sciences Program, UMDNJ-Graduate School of Biomedical Sci.)
Edward Garay (Biomedical Sciences Program, UMDNJ-Graduate School of Biomedical Sci.)
Ethan Fitzpatrick (Biomedical Sciences Program, UMDNJ-Graduate School of Biomedical Sci.)
Ahmet Tonceroglu (Biomedical Sciences Program, UMDNJ-Graduate School of Biomedical Sci.)
Xiangwen Chen (Department of Laboratory Medicine and Pathology, UMDNJ-Graduate School

Crystal DiCosmo (Department of Laboratory Medicine and Pathology, UMDNJ-Graduate School of Biomedical Sciences)

TRAINING SUMMER AND ROTATING STUDENTS

of Biomedical Sciences)

- 1994 Karen Finnigan
 - won first prize in Summer Student Cancer Research Symposium
 - won The Research Award for Scientific Excellence at the New Jersey State Commission on Cancer Research Annual Research Symposium, 1995
- 1994 Paul Maloof
- 1995 John Chung
- 1995 Daniel Fulop
- 1996 Christine Torigian
- 1996 Michelle A. Fanale
 - first prize, Summer Student Cancer Research Symposium
 - second prize, Annual NJMS Summer Student Res. Symp. (highest allowed)
 - first prize, First UMDNJ Statewide Med. Student Resch. Competition, 1997
 - Went on to Medical Oncology Fellowship at MD Anderson Cancer Center
- 1996 Annie Lin
- 1997 Mark Solomon
- 1997 Myrna S. Uytingco
 - won second prize in Summer Student Cancer Research Symposium
- 1997 Joseph Golowa
- 1997 Renato Apolito

- won first prize in NJMS Summer Student Cancer Research Symposium
- 1999 Lydia Choi
 - selected as 1 of 40 students nationally for oral presentation at The National Student Research Forum, University of Texas Medical Branch, Galveston, TX
 - won The National Student Research Forum Oncologic Research Award,
 - 2000 won the Gallo Research Award for Scientific Excellence, 2000 Annual Retreat on Cancer Research in New Jersey, CINJ and the NJCCR
 - 2004 Went on to Cancer Research Fellowship at Sloan Kettering
- 1999 Elizabeth Scheff
 - won first prize in NJMS Summer Student Cancer Research Symposium
 - won first prize in NJMS Summer Student Research Symposium
 - selected as 1 of 40 students nationally for oral presentation at The National Student Research Forum, University of Texas Medical Branch, Galveston, TX
- 2000 Ilan Seth Weinberg

Jeaudine Evadne Bontemps

Mateusz Opyrchal (MD/PhD rotation)

Jason Solomon

- 2001 Wei Bu (PhD rotation)
- 2001 Judith Barrios (PhD rotation)
- 2002 Michael Lindy won Award for Scientific Excellence at the CINJ-New Jersey State Commission on Cancer Research Annual Retreat, 2002
- 2002 Ankoor Shah
 - won travel award to National Meeting in NJMS Summer Student Cancer Research Symposium
- 2003 Vineetha Joseph
 - won book award in NJMS Summer Student Cancer Research Symposium
- 2003 Michael Lindy -
 - won travel award to National Meeting in NJMS Summer Student Cancer Research Symposium
 - selected as 1 of 40 students nationally for oral presentation at The National Student Research Forum, Univ. of Texas Medical Branch, Galveston, TX
- 2004 Mark Solomon
 - won book award in NJMS Summer Student Cancer Research Symposium
- 2004 Bryan Benn MD/PhD student
 - won book award in NJMS Summer Student Cancer Research Symposium
- 2005 Ethan Fitzpatrick (PhD rotation)
- 2005 Sylvia Vasquez
- 2006 Joanna Sesti
- 2006 Ahmet Tunceroglu (MD/PhD rotation)
- 2006 Aaron Rockoff (with Dr. Christakos, Biochemistry)
- 2007 Tara Tendler

TRAINING POST-DOCTORAL FELLOWS

1995-2000 Qin Wang, MD

won The New Jersey Research Award for Scientific Excellence (Smith-Kline Beecham Oncology, 1997

 won New Jersey Cancer Commission Outstanding Breast Cancer Research Post-Doctoral Fellowship, 1997

won Gallo Research Award for Scientific Excellence at the New Jersey
 State Commission on Cancer Research Annual Retreat, 1999, 2000

2001-2002 Petra Archibald, PhD

 won The Gallo Research Award for Scientific Excellence at the New Jersey State Commission on Cancer Research Annual Retreat, 2001

- won the 2001 American Association for Cancer Research Minority

Scholars in Cancer Research Award

 won the 2001 American Association for Cancer Research-Inglenook Vineyards Scholar-in-Training Award

2001-2002 Rachna Chandra, PhD 2003 Monika Boots, PhD

TRAINING GRADUATE STUDENTS

2002-2009 Judith Barrios (PhD)

2007-2008 Christopher K. Hansen (MS)

TRAINING FACULTY

2004-2006 Mentor for ASCO Young Investigator Award to Eric Cohen, MD, Assistant Professor, Dept. of Surgery, UMDNJ-NJMS

PROFESSIONAL ACTIVITIES

- 1988 Speaker: ZWO/TNO/NIH Symposium on Factors and Vectors in Hemopoiesis, The Hague, The Netherlands. Title: Use of retrovirally-mediated gene transfer for gene therapy in ADA deficiency.
- 1992 Speaker: American Society of Clinical Oncology, San Diego, CA. Title: Retroviral gene transfer of the hbFGF gene in human stroma.
- 1992 Speaker: International Society of Hematology, Providence, RI. Title: Cycle-activation of high proliferative potential cells (HPPC) in mice administered high doses of cytosine arabinoside (Ara-C).
- 1994 Speaker: American Society of Clinical Oncology, Dallas, TX. Title: MCF7 human breast cancer cells are negatively regulated by overexpression of basic fibroblast growth factor (bFGF).
- 1994-2006 Speaker, approx. 2 times/year UMDNJ-University Hospital Tumor Conference
- 1995 Speaker: Fourth International Conference on Gene Therapy of Cancer, San Diego, Ca. Title: Overexpression of retrovirally transduced basic FGF in MCF-7 human breast cancer cells downregulates Bcl-2 and sensitizes cells to chemotherapy-induced apoptosis.
- 1996 Speaker: First Annual New Jersey Breast Cancer Research Symposium, Rider University, Lawrenceville, NJ. Title: Basic FGF causes growth arrest in MCF-7 human breast cancer cells while inducing both mitogenic and inhibitory G₁ events.
- 1997 Speaker: Second Annual New Jersey Breast Cancer Research Symposium, UMDNJ-Robert Wood Johnson Medical School, New Brunswick, NJ. Title: 1,25-Dihidroxyvitamin D₃ and all-*trans* retinoic acid sensitize breast cancer cells to the effects of chemotherapeutic agents.
- 1998 Speaker at Cancer Institute of New Jersey Protocol Advisory Committee meeting

- 1998 Plenary Session Speaker at The Annual Retreat on Cancer Research in New Jersey
- 1999 Session Speaker at The Annual Retreat on Cancer Research in New Jersey
- 1999 Plenary Session Speaker, The Third New Jersey Breast Cancer Research Symposium
- 1999 Speaker, ACS/Cancer Care/CINJ/NJCCR/NJHD/St. Barnabas Hosp. Conference on Critical Decisions in Cancer for the 21st Century, Iselin NJ,
- 2000 Plenary Session Speaker, The Department of Defense Breast Cancer Research Program Meeting, "Era of Hope". Atlanta, GA, June 2000
- 2001 Speaker at Cancer Institute of New Jersey Protocol Advisory Committee meeting
- 2001 Speaker at the New Jersey State Commission on Cancer Research Symposium, "Sharing Perspectives on Cancer Research: Cancer Researchers Reach Out", Rider University, Lawrenceville, NJ
- 2002 Session Chair, Transcriptional regulation and oncogenensis/molecular mechanisms of tumor growth. Annual Retreat on Cancer Research in NJ, The Cancer Institute of NJ and the NJ State Commission on Cancer Research.
- 2006 Session Chair and Speaker, 10th Anniversary of the New Jersey Breast Cancer Research Fund Symposium

Seminars Given:

- 1993 Division of Hematologic Oncology, Memorial Sloan-Kettering Cancer Center, NY
- 1993 Department of Microbiology and Molecular Genetics, UMDNJ-NJ Medical School
- 1994 Department of Medicine Research Seminar, UMDNJ-NJ Medical School
- 1994 Hematology/Oncology Grand Rounds, East Orange Veterans Administration Hospital
- 1995 Hematology/Oncology Grand Rounds, New York Medical College
- 1995 Department of Laboratory Medicine and Pathology, UMDNJ-NJ Medical School
- 1996 The Center for Laboratory Investigation, UMDNJ-NJ Medical School
- 1996 Department of Surgery Research Conference, UMDNJ-NJ Medical School
- 1996 Department of Surgery Grand Rounds, UMDNJ-New Jersey Medical School
- 1997 Speaker, UMDNJ-NJMS Summer Student Cancer Research Symposium
- 1997 Department of Medicine Research Seminar, UMDNJ-NJ Medical School
- 1998 Hematology/Oncology Grand Rounds, East Orange Veterans Administration Hospital
- 1998 Keynote Address, UMDNJ-NJMS Summer Student Cancer Research Symposium
- 1998 Department of Obstetrics and Gynecology Grand Rounds, UMDNJ-NJ Med. School
- 1999 Department of Obstetrics and Gynecology Resident Conference, UMDNJ-NJMS
- 1999 Department of Medicine Grand Rounds, East Orange Veterans Administration Hosp.
- 2000 Speaker, UMDNJ-NJMS Summer Student Cancer Research Symposium
- 2000 Department of Medicine Grand Rounds, UMDNJ-New Jersey Medical School
- 2000 Department of Microbiology and Molecular Genetics, UMDNJ-NJ Medical School
- 2001 Department of Obstetrics and Gynecology Resident Conference, UMDNJ-NJMS
- 2001 Department of Anatomy, Cell Biology & Injury Sciences Res. Seminar UMDNJ-NJMS
- 2001 Speaker at Cancer Institute of New Jersey SPORE project seminar series
- 2001 Speaker, UMDNJ-NJMS Summer Student Cancer Research Symposium
- 2001 Division of Urology, Dept. of Surgery Grand Rounds, UMDNJ-NJ Med. School
- 2001 Department of Medicine Research Seminar, UMDNJ-New Jersey Medical School
- 2002 Division of Endocrinology, Dept. Medicine Research Seminar, UMDNJ-NJMS
- 2002 Speaker, UMDNJ-NJMS Summer Student Cancer Research Symposium
- 2003 Speaker, UMDNJ-NJMS Summer Student Cancer Research Symposium

- 2003 Speaker at Cancer Institute of New Jersey breast cancer seminar series
- 2003 Seminar speaker at Rider University Department of Biology
- 2004 Research seminar, UMDNJ-University Hospital Tumor Board
- 2004 Speaker, UMDNJ-NJMS MD/PhD Student Research Seminar
- 2004 Speaker, UMDNJ-NJMS Summer Student Cancer Research Symposium
- 2004 Speaker Governor's School Students Cancer Biology Course
- 2005 Speaker, UMDNJ-NJMS Biomedical Sciences Program Research Seminar Series
- 2005 Seminar speaker at Rider University Department of Biology
- 2005 Speaker, Hematology/Oncology Research Conference, East Orange VA Hospital
- 2005 Speaker, Division of Endocrinology Research Conference, UMDNJ-NJMS
- 2006 Speaker, Research Conference, Department of Medicine, UMDNJ-NJMS
- 2006 Overview of Paradigms for Clinical Research, UMDNJ-University Hospital Cancer Center Grand Rounds
- 2006 Speaker, Division of Endocrinology Research Conference, UMDNJ-NJMS
- 2007 Speaker, UMDNJ-NJMS MD/PhD Student Research Seminar
- 2007 Seminar speaker, Department of Biochemistry and Molecular Biology, UMDNJ-New Jersey Medical School
- 2007 Speaker, UMDNJ-NJMS Summer Student Cancer Research Symposium
- 2007 Speaker, UMDNJ-NJMS Summer Student Cancer Research Seminar Series
- 2008 Speaker, UMDNJ-NJMS Mini Med School
- 2008 Speaker, Cancer Institute of New Jersey Breast Cancer Program Seminar Series
- 2008 Speaker, UMDNJ-NJMS Summer Student Cancer Research Symposium
- 2008 Speaker, McNair Summer Student Scholars Program, Rider University, Lawrenceville, NJ
- 2008 Speaker, UMDNJ-NJMS Premed Honors Program
- 2009 Speaker, UMDNJ-NJMS Summer Student Cancer Research Symposium

Meeting Organizer:

1996 Co-chair: First New Jersey Breast Cancer Research Symposium Co-chair: Third New Jersey Breast Cancer Research Symposium

Manuscript reviewer:

Biochemical Pharmacology

Biochimica et Biophysica Acta

British Journal of Cancer

Cancer

Cancer Letters

Cancer Research

Clinical Cancer Research

Endocrinology

Experimental Cell Research

FEBS Letters

Journal of Biological Chemistry

Journal of Cellular Biochemistry

Journal of Cellular Physiology

Molecular Cancer Therapeutics Molecular Pharmacology Oncogene Pharmacological Research

Grant reviewer

ant	reviewer	
	1994-2000	Foundation of UMDNJ
	2000	Dutch Cancer Society (ad hoc)
	2001	National Science Foundation (ad hoc)
	2003	Foundation for Science and Technology, Portugal Ministry of Science
	2004	Department of Defense Breast Cancer Research Program
		ENDO-2 Study Section
		EPI-Adhoc Study Section, Chair
	2005	Susan G. Komen Breast Cancer Foundation Tumor Cell Biology I Study
		Section
	2005	National Cancer Institute, NIH Cancer Epidemiology/Cancer Prevention
		Small Grants Study Section
	2006	Susan G. Komen Breast Cancer Foundation Tumor Cell Biology V Study
		Section
	2006	Department of Defense Breast Cancer Research Program Molecular
		Biology & Genetics Peer Review Panel III
	2007	Susan G. Komen Breast Cancer Foundation Tumor Cell Biology I Study
		Section
	2007	California Breast Cancer Research Program Tumor Progression
		Review Committee
	2007	National Science Foundation (ad hoc)
	2007	National Cancer Institute, Innovative Technologies for Molecular
		Analysis of Cancer (IMAT) review panel
	2007	Department of Defense Breast Cancer Research Program Molecular
		Biology & Genetics Peer Review Panel III
	2008	Susan G. Komen Breast Cancer Foundation Tumor Cell Biology I Study
	0000	Section
	2008	National Cancer Institute, Innovative Technologies for Molecular
	0000	Analysis of Cancer (IMAT) review panel
	2008	California Breast Cancer Research Program Tumor Progression
	2000	Committee
	2008	Israeli Science Foundation (ad hoc)
	2008	Department of Defense Breast Cancer Research Program Molecular
	2009	Biology & Genetics Peer Review Panel II
	2009	Chair, DOD Breast Cancer Research Program Concept Grant Molecular
	2009	Biology & Genetics Peer Review Panel I Department of Defense Breast Cancer Research Program Molecular
	2009	Biology & Genetics Peer Review Panel II
	2009	Foundation of UMDNJ Biomedical Research Support Program
	2009	Department of Defense Breast Cancer Research Program Molecular
		Biology & Genetics Peer Review Panel II
		Diology & Dollottoo Foot Neview Fallet II

17. PRINCIPAL CLINICAL AND HOSPITAL SERVICE RESPONSIBILITIES

Weekly Oncology clinic, UMDNJ- University Hospital Service Attending two months a year in Oncology, UMDNJ-University Hospital

18. MAJOR ADMINISTRATIVE RESPONSIBILITIES

1993-present Director, Laboratory of Molecular Oncology

1996-1999 Interim Director, Division of Oncology, Department of Medicine. UMDNJ-New Jersev Medical School

1999-2001 Interim Director, Division of Oncology/Hematology, Department of Medicine, UMDNJ-New Jersey Medical School

1998-2001, 2006-present Chair, NJ Commission on Cancer Research Breast Cancer Advisory Group

1998-2001 Associate Director, Clinical Research, UMDNJ-NJMS Cancer Center

1998-2003 Associate Director (clinical), P.D./PhD Program, UMDNJ-Graduate School of Biomedical Sciences

2005-present Director, Clinical Research Office, UMDNJ-NJ Medical School/ University Hospital Cancer Center

2009-present Co-Medical Director, Center for Clinical and Translational Science, UMDNJ-NJ Medical School

19. PRIVATE PRACTICE

None

BIBLIOGRAPHY

20. ARTICLES

- 1. Jaynes EN, Grant PG, Giangrande G, **Wieder R**, Cooperman BS. (1978) Photoinduced affinity labeling of the <u>Escherichia coli</u> ribosome puromycin site. Biochemistry 17: 561-569.
- 2. **Wieder R**, Wetmur JG. (1981) One hundred-fold acceleration of DNA renaturation rates in solution. Biopolymers 20: 1537-1547.
- 3. **Wieder R**, Wetmur JG. (1982) Factors affecting the kinetics of DNA reassociation in phenol-water emulsion at high DNA concentrations. Biopolymers 21: 665-677.
- 4. **Wieder R**. (1982) Techniques for accelerating DNA renaturation and their preliminary application to gene isolation methods. Thesis. The City University of New York.
- 5. Cornetta K, **Wieder R**, Anderson WF. (1989) Gene transfer into primates and prospects for gene therapy in humans. Progress in Nucleic Acids Research and Molecular Biology 36:311-322.
- 6. **Wieder R**, Cornetta K, Kessler SW, Anderson WF. (1991) Increased efficiency of retroviral-mediated gene transfer and expression in primate bone marrow progenitors following 5-FU-induced hematopoietic suppression and recovery. Blood 77: 448-455.
- 7. Wieder R. (1991) Cryopreserved primate bone marrow cells can be used for retroviral-

mediated gene transfer. Human Gene Therapy 2: 323-326.

- 8. **Wieder R**, Barak V, Ben-Ishay Z. (1995) High-efficiency retroviral gene transfer into murine high-proliferative-potential cells cycle-activated by cytosine arabinoside. Human Gene Therapy 6: 865-871.
- 9. Menzel T, Rahman Z, Calleja E, White K, Wilson EL, **Wieder R**, Gabrilove J. (1996) Elevated intracellular level of basic fibroblast growth factor correlates with stage of chronic lymphocytic leukemia and is associated with resistance to fludarabine. Blood 87: 1056-1063. 10. Fenig E. **Wieder R**, Paglin S, Wang H, Persaud R, Haimovitz-Friedman A, Fuks Z, Yahalom J. (1997) Basic fibroblast growth factor confers growth inhibition and Mitogenactivated Protein Kinase activation in human breast cancer cells. Clinical Cancer Research 3: 135-142.
- 11. Wang H, Rubin M, Fenig E, DeBlasio T, Mendelsohn J, Yahalom J and **Wieder R**. (1997) Basic FGF causes growth arrest in MCF-7 human breast cancer cells while inducing both mitogenic and inhibitory G₁ events. Cancer Research 57: 1750-1757.
- 12. **Wieder R**, Wang H, Shirke S, Wang Q, Menzel T, Feirt N, Jakubowski AA and Gabrilove JL. (1997) Low level expression of basic FGF upregulates Bcl-2 and delays apoptosis, but high intracellular levels are required to induce transformation in NIH 3T3 cells. Growth Factors 15:41-60.
- 13. **Wieder R**, Fenig E, Wang H, Wang Q, Paglin S, Menzel T, Gabrilove J, Fuks Z, Yahalom J. (1998) Overexpression of basic fibroblast growth factor in MCF-7 human breast cancer cells: lack of correlation between inhibition of cell growth and MAP kinase activation. J. Cellular Physiology 177:411-425.
- 14. Wang Q, Maloof P, Wang H, Fenig E, Stein D, Nichols G, Denny TN, Yahalom J and **Wieder R**. (1998) Basic fibroblast growth factor (bFGF) downregulates Bcl-2 and promotes apoptosis in MCF-7 human breast cancer cells. Experimental Cell Research 238:177-187.
- 15. Maloof P, Wang Q, Wang H, Stein D, Denny TN, Yahalom J, Fenig E and **Wieder R**. (1999) Overexpression of retrovirally transduced basic FGF in MCF-7 human breast cancer cells downregulates Bcl-2 and sensitizes cells to chemotherapy-induced apoptosis. Breast Cancer Research and Treatment 56:153-167.
- 16. Fenig E, Livnat T, Sharkon-Polak S, Wasserman L, Beery E, Lilling G, Yahalom J, **Wieder, R**, Nordenberg J. (1999) Basic fibroblast growth factor potentiates cisplatinum-induced cytotoxicity in MCF-7 human breast cancer cells. J. Cancer Res. Clin. Onc. 125:556-562.
- 17. Korah R, Sysounthone V, Golowa Y, and **Wieder R**. (2000) Basic fibroblast growth factor confers a more differentiated phenotype in MDA-MB-231 human breast cancer cells. Cancer Research 60:733-740.
- 18. Wang Q, Yang W, Uytingco MS, Christakos S and **Wieder R**. (2000) 1,25(OH)₂ vitamin D₃ and all-*trans* retinoic acid sensitize breast cancer cells to chemotherapy-induced cell death.

Cancer Research. 60:2040-2048.

- 19. Korah R, Sysounthone V, Scheff E, and **Wieder R**. (2000) Intracellular FGF-2 promotes differentiation in T47-D breast cancer cells. Biochem. Biophys. Res. Comm. 277:255-260.
- 20. Wang Q, Lee D, Sysounthone V, Chandraratna RAS, Christakos S, Korah R, and **Wieder** R. (2001) 1,25-dihydroxyvitamin D₃ and retinoic acid analogues induce differentiation in breast cancer cells with function- and cell-specific additive effects. Breast Cancer Res. Treat. 67:157-168.
- 21. Fenig E, Kanfi Y, Wang Q, Beery E, Livnat T, Wasserman L, Lilling G, Yahalom J, **Wieder** R Nordenberg J. (2001) Role of transforming growth factor beta in the growth inhibition of human breast cancer cells by basic fibroblast growth factor. Breast Cancer Res. Treat. 70: 27-37.
- 22. **Wieder R**, Pavlick AC, Bryan M, Hameed M, Baredes S, Pliner L, Saunders T and Korah R. (2002) Phase I/II trial of Accutane as a potentiator of carboplatin and Taxol in squamous cell carcinomas. American J. Clinical Oncology 25: 447-450.
- 23. **Wieder R**, Novick SC, Hollis BW, Bryan M, Chanel SM, Owusu K, Camastra D, Saunders T, Pliner L, Harrison J, Bonate P, Williams T, Soignet S. (2003) Pharmacokinetics and Safety of ILX23-7553, a Non-calcemic-Vitamin D₃ Analogue, in a Phase I Study of Patients with Advanced Malignancies. Investig. New Drugs 21: 445-452.
- 24. Wang Q and **Wieder R.** (2004) All-*trans* retinoic acid potentiates Taxotere-induced cell death mediated by jun N-terminal kinase in breast cancer cells. Oncogene 23: 426-433.
- 25. Korah R, Choi L, Barrios J and **Wieder R**. (2004) Constitutive expression of FGF-2 abrogates focal adhesion signaling in MDA-MB-231 breast cancer cells. Breast Cancer Research and Treatment 88: 17-28 (Erratum color photos (2005) 89: 319 322).
- 26. Korah R, Boots M, and **Wieder R**. (2004) Integrin $\alpha 5\beta 1$ promotes survival of growth-arrested breast cancer cells: an *in vitro* paradigm for breast cancer dormancy in bone marrow. Cancer Research 64: 4514-4522.
- 27. Najmi S, Korah R, Chandra R, Abdellatif M, **Wieder R**. (2005) Flavopiridol blocks integrin-mediated survival in dormant breast cancer cells. Clinical Cancer Research 11:2038-2046.
- 28. **Wieder R**. Insurgent micrometastases: sleeper cells and harboring the enemy. (2005) J. Surgical Oncology 89:207-210.
- 29. Fitzpatrick E, McBride S, Yavelow J, Najmi S, Zanzucchi P and **Wieder R**. Microfluidic techniques for single cell protein expression analysis. (2006) Clin. Chem. 52:1080-1088.
- 30. Korah R, Das K, Lindy ME, Hameed M and **Wieder R**. Co-ordinate loss of FGF-2 and laminin 5 expression during neoplastic progression of mammary duct epithelium. (2007)

Human Pathology 38:154-160.

- 31. Bryan M, De La Rosa N, Hill AM, Amadio WJ, **Wieder R**. (2008) Influence of prescription benefits on pain control in patients with cancer. Pain Medicine 9:1148–1157.
- 32. Dhawan P, **Wieder R**, Christakos S. (2009) CCAAT Enhancer Binding Protein Alpha is a Molecular Target of 1,25Dihydroxyvitamin D₃ in MCF-7 Breast Cancer Cells. J. Biol. Chem. 284:3086-3095.
- 33. Barrios, J and **Wieder R**. (2009) Dual FGF-2 and intergrin $\alpha 5\beta 1$ signaling mediate GRAF-induced RhoA inactivation in a model of breast cancer dormancy. Cancer Microenvironment (in press).

21. BOOKS, MONOGRAPHS AND CHAPTERS

- 1. **Wieder R**, Kessler SW, Wagemaker G, Anderson WF. (1988) Differential retroviral gene transfer into primate bone marrow precursors fractionated on an albumin gradient. In: Gale RP and Champlin R, eds., UCLA Symposia on Molecular and Cellular Biology, New Series, vol 91: Bone Marrow Transplantation: Current Controversies, Alan R. Liss, Inc, New York, pp 379-388.
- 2. **Wieder, R**. Selection of Methods for Measuring Proliferation, in Cell Growth, Differentiation and Senescence: A Practical Approach. G. Studzinski, ed. Oxford University Press, New York, NY, 1999, pp 1-32.
- 3. **Wieder, R**. TUNEL assay as a measure of chemotherapy-induced apoptosis. Methods in Molecular Medicine, vol. 111: Chemosensitivity: Vol. 2: In Vivo Models, Imaging, and Molecular Regulators. R.D Blumenthal, ed., Humana Press, Inc., Totowa, NJ, 2005, pp 43-54.

22. ABSTRACTS

- 1. **Wieder R**, Wetmur JG. (1981) Optimum methods for acceleration of DNA renaturation rates. Federation Proceedings 40: 1849.
- 2. Zwiebel JA, Kantoff PW, Eglitis MA, Kohn D, Muenchau D, McLachlin JR, Karson E, **Wieder R**, Yu S-F, Blaese MR, Gilboa E, Anderson WF. (1986) Gene transfer and expression using a family of retroviral vectors. Blood 68: 307a.
- 3. Cornetta K, Moen R, Gillio A, Culver K, **Wieder R**, Blaese RM, O'Reilly R, Anderson WF. (1988) Fate of murine helper virus in non-human primates. J Cellular Biochem, Supplement 12B.
- 4. **Wieder R**, Zwiebel JA, Wagemaker G, Anderson WF. (1988) Enhanced retroviral gene transfer into primate bone marrow progenitor cells enriched by discontinuous albumin

- gradients. J Cellular Biochem, Supplement 12C, K221.
- 5. **Wieder R**, Cornetta K, Kessler S, Anderson WF. (1988) Kinetics of 5-FU-induced bone marrow suppression and recovery: effects on the efficiency of retroviral gene transfer in non-human primates. Blood 72: 105a.
- 6. **Wieder R**, Cornetta K, Kessler S, Anderson WF. (1989) Improved efficiency of retroviral-mediated gene transfer and expression in primate hematopoietic progenitors following 5-FU-induced bone marrow suppression and recovery. J Cellular Biochem, Suppl. 13C, H229.
- 7. **Wieder R**, Shirke S, Kehagias E, Gilboa E, Rifkin DB, Wilson EL, Jakubowski AA, Gabrilove JL. (1991) Constitutive expression of retrovirally transduced basic FGF in NIH 3T3 cells causes phenotypic transformation and modulates hematopoiesis. Blood 78: 301a.
- 8. **Wieder R**, Shirke S, Kehagias E, Jakubowski A, Wilson EL, Gabrilove JL. (1992) <u>In vitro</u> stimulation of myelopoiesis by constitutive expression of basic FGF in retrovirally transduced NIH 3T3 cells. J Cellular Biochemistry, Supplement 16C, M446.
- 9. **Wieder R**, Shirke S, Kehagias E, Jakubowski AA, Wilson EL, Gabrilove JL. (1992) NIH 3T3 cells transduced with basic FGF stimulate myelopoiesis <u>in vitro</u>. J Cellular Biochemistry, Suppl. 16F, V229.
- 10. **Wieder R**, Shirke S, Wilson EL, Gabrilove JL. (1992) Retroviral Gene transfer of the human basic fibroblast growth factor (hbFGF) gene in human stroma. Cancer Biology and Molecular Genetics, 11:105, #241. (selected for oral presentation) (Winner of ASCO Travel Award)
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23. REPORTS

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EXHIBIT B

Figure 1. Nonrad GEArray Q series gene chip microarray analysis of MCF-7 cells incubated with and without FGF-2 for 5 days on tissue culture dishes coated with fibronectin

Α.

+ FGF-2 - FGF-2

Figure 1. Nonrad GEArray Q series Human Extracellular Matrix and Adhesion Protein chip (Super Array, Bethesda, MD) microarray analysis of MCF-7 cells incubated for 5 days on tissue culture dishes coated with fibronectin 20 μg with and without the presence of FGF-2 10 ng/ml. Approximately one third as many cells remained in the FGF-2-treated population as in the control cells. Arrows point to integrin αV (solid line, spot 43) and $\beta 6$ (dotted line, spot 50) mRNA's that are not expressed in the starting population and unchanged in the surviving population. Boxes are drawn around the control gene cDNAs on the two chips consisting of GAPDH, Cyclophyllin A, ribosomal L23 and β actin as positive controls and PUC18 plasmid DNA and blanks as negative controls.

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Array Layout Table with Gene Symbol and Position Information

Human Adhesion & Extracellular Matrix Molecules GEArray Q series version 1

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ADAMTS1	ADAMTS8	CASP8	CASP9	CAV1	CD44	CDH1	CEACAM5
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· CNTN1	COL18A1	COLIAI	COL4A2	CST3	CTNNA1	CTNNAL1	CTNNB1
9	10	11	12	13	14	15	16
CTNND1	CTNND2	CTSB	CTSD	CTSG	CTSL	DCC	ECM1
17	18	19	20	21	22	23	24
FG8	FN1	HPSE	ICAM1	ITGA1	ITGA10	ITGA11	ITGA2
25	26	27	28	29	30	31	32
ITGA2B	ITGA3	ITGA4	ITGA5	ITGA6	ITGA7	ITGA8	ITGA9
33	34	35	36	37	38	39	40
ITGAL	ITGAM	ITGAV	ITGAX	ITGB1	ITGB2	ITGB3	ITG84
41	42	43	44	45	46	47	48
ITG85	ITGB6	ITGB7	ITGBB	LAM81	LAMC1	MGEA5	MICA
49	50	51	52	53	54	55	56
MMP1	MMP10	MMP11	MMP12	MMP13	MMP14	MMP15	MMP16
57	58	59	60	61	62	63	64
MMP17	MMP2	MMP20	MMP24	MMP26	MMP3	MMP7	ММРВ
65	66	67	68	69	70	71	72
MMP9	NCAM1	NRCAM	PECAM1	PLAT -	PLAU	PLAUR	SELE
73	74	75	76	77	78	79	80
SELL	SELP	SERPINB2	SERPINBS	SERPINE1	SPARC	SPP1	THBS1
81	82	83	84	85	86	87	88
THBS2	THBS3	TIMP1	TIMP2	TIMP3	TMPRSS4	VCAM1	VTN
89	90	91	92	93	94	95	96
PUC18	PUC18	PUC18	Blank	Blank	Blank	GAPD	GAPD
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10. 51.34         Will (1921)         MCG-25.         Averagents accepted at the Companion accessed at the Companion and Companion and Companion at the Companio	4	14, 214962	103202	(SMC)	amine, gamma 1 (formatly (ALIG2)	amento 62
16.5159   Nil. 2002177   Nil.   Nich.   Nic.   Nich.   Nich.   Nich.   Nich.   Nich.   Nich.   Nich.   Nich.	55	Hs 5/34	-	MGEAS	Meningioma expressed entigen 5 (hyalumandase)	Hydiuromdases
1, 17.253   10.003.13   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.003.14   10.0	3	H 80308		NC.	tomo sapiens MICA gene, allete MLIC-18	MUC:18
11.7255   343 (202155   348PT)   Marin metalopolements 12 (Incomplate elistate)     14.15524   A.27166   A.2716   A.27166   A.2716   A.27166   A.2716   A.27166   A.	2	11, 63109	X05231	MMP1	Massix metallogrotemasa 1 (intercitual codlagenasa)	companda .
11. 155.224         3.27106         3.42(71)         Aut/P11	\$6	145 7253	Nu 002425	OLCHIN	Matrix metaboprobenuse 10 (strometysin 2)	stametysin 2
17.555   27800   131912   441914   4144916   4164916   416491   415750   418751   444914   4144916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   4164916   416	55	H. 155.124	A57786	MAP 11	Human strometysm-3	Show shain.
14.3796   X75.00	8	1555	123906	ZI divin	Matru metallopiotonase 12 (ma.rophage elastase)	mail opinatie
13.250   Vi 23.01   Map 14   Map 14   Map 15			10000	1,00,4	Little man alternate 13 frontstate 31	Collegenson 3
18.299   10.003.12   Augle   August	5	15 2936	K/Saug		Settling in the feet of the settle of the se	MT1-MMP
1, 10, 10, 10, 10, 10, 10, 10, 10, 10,	3	14 7399	715920		1. September 10. Stationary 2.	MAN 15
1, 1500   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0316   10.0	8	F E0343	Dealsh		Marine interestable of the marine and the marine an	MadP 16
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1373   13, 106171   44078   Main metalogueterase 21 (national continuos directors)   13, 1373   13, 106171   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078   14078	8	Ē.1.15	012200	2	pastro metalloprotembre 4 (generales A. 1210 personal.	
1, 17, 13   13   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100   100	3	129737	124 004771	02.dW	Matrix metanopiciemase 20 (enamelytin)	enamely sin
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1, 63.20   20233   13493   Main's matalogousman a 3 (stromegran )   projectionare)     14. 12.56   207819   Main's matalogousman a 3 (stromegran)   projectionare     14. 12.56   20.001242   Main's matalogousman a 1 (stational a 10.0012)     14. 12.102   Main's matalogousman a 1 (stational a 10.0012)     14. 12.102   Main's matalogousman a 1 (stational a 10.0012)     14. 12.102   Main's matalogousman a 1 (stational a 10.0012)     15. 12.102   Main's matalogousman a 1 (stational a 10.0012)     16. 12.102   Main's matalogousman a 1 (stational a 10.0012)     16. 12.102   Main's matalogousman a 1 (stational a 10.0012)     16. 12.102   Main's matalogousman a 1 (stational a 10.0012)     16. 12.102   Main's matalogousman a 1 (stational a 10.0012)     16. 12.102   Main's matalogousman a 1 (stational a 10.0012)     16. 12.102   Main's matalogousman a 1 (stational a 10.0012)     16. 12.102   Main's matalogousman a 1 (stational a 10.0012)     17. 12.102   Main's matalogousman a 1 (stational a 10.0012)     18. 12.102   Main's matalogousman a 1 (stational a 10.0012)     18. 12.102   Main's matalogousman a 1 (stational a 10.0012)     18. 12.102   Main's matalogousman a 1 (stational a 10.0012)     18. 12.102   Main's matalogousman a 1 (stational a 10.0012)     18. 12.102   Main's matalogousman a 1 (stational a 10.0012)     18. 12.102   Main's matalogousman a 1 (stational a 10.0012)     18. 12.102   Main's matalogousman a 1 (stational a 10.0012)     18. 12.102   Main's matalogousman a 1 (stational a 10.0012)     18. 12.102   Main's matalogousman a 1 (stational a 10.0012)     18. 12.102   Main's matalogousman a 1 (stational a 10.0012)     18. 12.102   Main's matalogousman a 1 (stational a 10.0012)     18. 12.102   Main's matalogousman a 1 (stational a 10.0012)     18. 12.102   Main's matalogousman a 1 (stational a 10.0012)     18. 12.102   Main's matalogousman a 1 (stational a 10.0012)     18. 12.102   Main's matalogousman a 1 (stational a 10.0012)     18. 12.102   Main's matalogousman a 1 (stational a 10.0012)     18. 12.102	6	15 204732	÷	SCGN1	Homo capares mairix meladopcitemase-26 (MMP26) mRNA	MAPZO
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11. 1517.13 (1902) 2. 124. 125. 125. 125. 125. 125. 125. 125. 125	1	11: 22:56	\$37819	MAPT	Matrix matalloprotesties 7 (matriysin, identie)	Thethy'sin,
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13   15   15   15   15   15   15   15	22	Hs. 73962	JIM_002424	9	אשונוא נואנישונסלטניגטינישורים כן נואפיניניליאיש רבייניקאיים	Collegenase
11/19/82   M3241   M2A41   Maria & sistentian mosecule   Maria & sistentian mosecule   Maria & sistentian mosecule   Maria & sistentian mosecule   Maria & sistentian   Maria & sistentian mosecule   Maria & sistentian	12	15, 1517.38	-,	Вихи	Matrix metalloproteinese & (getatinase B. BZLD) getafinase, GZLD type IV	geleinate B
14   17   18   18   18   18   18   18   18			-	IV ANA	Coffee 6 adhering molecule 1	15
17.74(4)   Na. 003447   FECAM  Purms separa pataestercanheid cell santu-un molecula (CD3) eragen)   17.74(4)   Na. 003450   74.77   Picchael pataestercanheid cell santu-un molecula (CD3) eragen   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)   17.75(4)		500	4	MACAM	nausonal ced adhesam mostule	NRCAM
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No. 17214         DO0344         P.A.J.         Personage netheral contents           No. 17214         DO0344         P.A.J.         Personage netheral contents           No. 17260         III. (2075:0)         F.L.K.II.         American ordinate lead to personament of the content of	17	HS 274404	_	J.A.	Home express plasmingen activator, tissue (PLA18)	
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Gene name	TIME	TIMP?	TIMP3	TUPRSSA	VCAMI	virgnectes	- Piccia	HIC IS	PUCIE			0	GAFON	GAPOH	Lyclophen A	:)-claphilin .	Cyclophin A	Cyclophin A	AN SIA	RP, 134	p-refin	o.scta
Description	Tissue inhibitor of metalloproteinase 1 (exphred potentisting activity, collegenase inhibitor).	1 Issue mhatar of metallogratemase 2	Insue mhibitor of metallogrammes a 3 (Soriby fundes dystrophy, pseudomnemmetry)	Transmonbrane protesse, sering 4	vescular cell adhesion molecuia 1	Vitanetin (serum spreading factor, sometomental), complement 5.	PUCIS Plasmid CitA	PUC18 Plasmid DNA	PLC 18 Playmid Dita	Bank	#UP	Blank	Chyceraldehyde J. phosphale defydrogenase	Chceraidehyde-3-phosphata dehydrogenase	Homo septems peptidybroth isomasse A (cyclopidin A) (PPIA)	formo septems peptidyleproly/ isomerase A (cyclophian A) (PPIA)	Home septens presidy brane as a (cyclophin A) (PPIA)	Home sapiens peptidyforctyl isomerase A (cyclophilin A) (PPIA)	Ricosomal protein L13a(23 Kda hghly basic protein)	Piposomel protein (13a(23 Kds highly basic protein)	Bata Activ	Beta Activ
Symbol	Ida	11002	TIMPS	*NPRSS4	VCAM	3	NC III	5	PUCTE	Blenk	Bank	Hank	0.83	Odeo	¥lød	¥Jd.	₽H4	¥ldd	API 13A	RPC13A	ACT0	AC.18
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